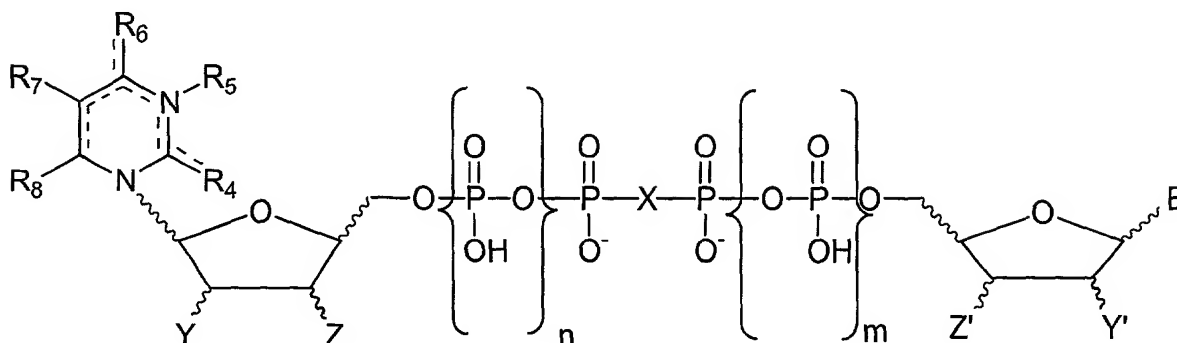


WHAT IS CLAIMED IS:

1. A compound of Formula IIIA:

Formula IIIA



wherein:

X is oxygen, methylene, difluoromethylene, imido;

n = 0, 1, or 2;

m = 0, 1, or 2;

n + m = 0, 1, 2, 3, or 4;

B is a purine or a pyrimidine residue linked through the 9- or 1-position, respectively;

Z = OH or N₃;

Z' = OH or N₃;

Y = H or OH;

Y' = H or OH;

provided that when Z is N₃, Y is H or when Z' is N₃, Y' is H;

R₄ is oxo, amino, cyano, aralkoxy, C₁₋₆ alkoxy, C₁₋₆ alkylamino, or dialkylamino;

R₅ is hydrogen, acyl or benzoyl, C₁₋₆ alkyl, phenyloxy, C₁₋₅ alkanoyl or

absent;

R₆ is oxo, hydroxy, mercapto, C₁₋₄alkoxy, C₇₋₁₂arylalkoxy, C₁₋₆alkylthio, amino, C₁₋₅ disubstituted amino, triazolyl, C₁₋₆alkylamino or di-C₁₋₄alkylamino, where the alkyl groups is optionally linked to form a heterocycle or link to N³ to form a substituted ring; or

R₅ and R₆ taken together form a 5-membered fused imidazole ring between positions 3 and 4 of the pyrimidine ring, which is optionally substituted on the 4- or 5- positions of the etheno moiety with C₁₋₄alkyl, phenyl, or phenyloxy, which themselves are optionally substituted;

R₇ is hydrogen, hydroxy, cyano, nitro, C₂₋₈alkenyl, C₁₋₄alkyl, phenyl, substituted C₂₋₈alkynyl, halogen, C₁₋₄alkyl, substituted C₁₋₄alkyl, CF₃, C₂₋₆ alkyl, C₂₋₃ alkenyl, allylamino, bromvinyl, ethyl propenoate, propenoic acid, C₂₋₃ alkynyl, substituted C₂₋₃alkynyl; or

R₆ and R₇ taken together form a 5 or 6-membered saturated or unsaturated ring bonded through N or O at R₆, such ring optionally contain substituents that themselves contain functionalities;

R₈ is hydrogen, amino or di-C₁₋₄alkylamino, C₁₋₄alkoxy, C₇₋₁₂arylalkoxy, C₁₋₄alkylthio, C₇₋₁₂arylalkylthio, carboxamidomethyl, carboxymethyl, methoxy, methylthio, phenoxy or phenylthio; provided that when R₈ is amino or substituted amino, R₇ is hydrogen;

provided that when B = adenine, adenine 1-oxide, or 1,N⁶-ethenoadenine, then:

(a) R₆ ≠ oxo when R₄ = oxo, Y = Z = OH and R₅ = R₇ = R₈ = H;

(b) R₇ ≠ Br when R₄ = R₆ = oxo, Y = Z = OH, and R₅ = R₈ = H;

provided that when B = adenine, then:

(a) R₆ ≠ amino when R₄ = oxo, Y = Z = OH, R₅ is absent, R₇ = R₈ = H, and n + m = 0, 1, or 2;

(b) R₇ ≠ CH₃ when R₄ = R₆ = oxo, Y = H, Z = OH, and R₅ = R₈ = H;

(c) R₇ ≠ F when R₄ = R₆ = oxo, Y = H, Z = OH, R₅ = R₈ = H and n + m = 2;

provided that when B = thymine, Y' = H and Z' = N₃; then R₇ ≠ F, when R₄ = R₆ = oxo, Y = OH, Z = OH, R₅ = R₈ = H, and n + m = 0;

provided that when B = thymine, Y' = H and Z' = N₃; then R₇ ≠ CH₃ when R₄ = R₆ = oxo, Y = H, Z = N₃, R₅ = R₈ = H, and n + m = 0;

provided that when B = guanine, then:

(a) R₆ ≠ oxo when R₄ = oxo, Y = Z = OH, R₅ = R₇ = R₈ = H and n + m = 1 or 2;

- (b) $R_6 \neq \text{amino}$ when $R_4 = \text{oxo}$, $Y = Z = \text{OH}$, R_5 is absent, $R_7 = R_8 = \text{H}$, $n+m=1$ or 2;

provided that when B is uridine, or 5-Br-uridine, then

- 5 (a) $R_6 \neq \text{oxo}$ when $R_4 = \text{oxo}$, $Y = Z = \text{OH}$ and $R_6 = R_7 = R_8 = \text{H}$;
 (b) $R_7 \neq \text{Br}$ when $R_4 = R_6 = \text{oxo}$, $Y = Z = \text{OH}$, and $R_5 = R_8 = \text{H}$;

provided that when B is 5-FU, then $R_7 \neq \text{F}$, when $R_4 = R_6 = \text{oxo}$, $Y = \text{H}$, $Z = \text{OH}$, $R_5 = R_8 = \text{H}$, and $n + m = 0$;

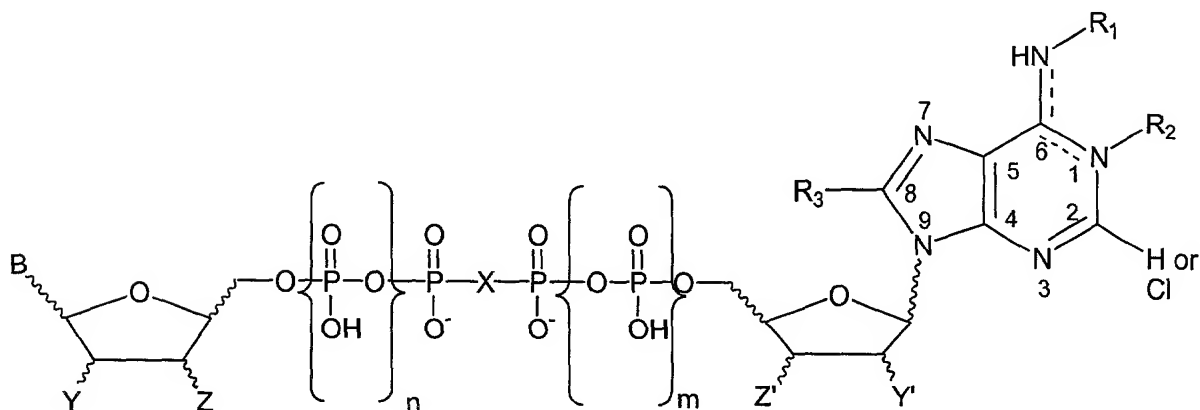
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provided that when B is cytosine, then $R_6 \neq \text{amino}$, when $R_4 = \text{oxo}$, $Y = Z = \text{OH}$, R_5 is absent, $R_7 = R_8 = \text{H}$, and $n + m = 1$, or 2; and

provided that when B is cytosine, then $R_6 \neq \text{oxo}$, when $R_4 = \text{oxo}$, $Y = Z = \text{OH}$ and $R_6 = R_7 = R_8 = \text{H}$, and $n + m = 2$.

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2. A compound according to Formula IIA:



Formula IIA

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wherein:

X is oxygen, methylene, difluoromethylene, imido;

n = 0, 1, or 2;

m = 0, 1, or 2;

n + m = 0, 1, 2, 3, or 4;

B is a purine residue linked through the 9- position;

Z = OH or N₃;

Z' = OH or N₃;

Y = H or OH;

Y' = H or OH;

provided that when Z is N₃, Y is H or when Z' is N₃, Y' is H;

R₁ is H, C₁₋₈alkyl, phenyl or phenyloxy, optionally substituted with halogen, hydroxy, C₁₋₄alkoxy, C₁₋₄alkyl, C₆₋₁₀aryl, carboxy, cyano, nitro, sulfonamido, sulfonate, phosphate, sulfonic acid, amino or substituted amino, wherein the amino is singly or doubly substituted by a C₁₋₄ alkyl and when doubly substituted, the alkyl groups are optionally linked to form a heterocycle; or A(C₁₋₆alkyl)CONH(C₁₋₆alkyl)B wherein A and B are amino, mercapto, hydroxy or carboxyl;

R₂ is O or is absent; or

R₁ and R₂ taken together forms a 5-membered fused imidazole ring, which is optionally substituted on the 4- or 5- positions of the etheno moiety with C₁₋₄alkyl, phenyl or phenyloxy, optionally substituted with halogen, hydroxy, C₁₋₄alkoxy, C₁₋₄alkyl, C₆₋₁₀aryl, arylalkyl, carboxy, cyano, nitro, sulfonamido, sulfonate, phosphate, sulfonic acid, amino or substituted amino, wherein the amino is singly or doubly substituted by a C₁₋₄ alkyl and when doubly substituted, the alkyl groups is optionally linked to form a heterocycle; and

R₃ is H, C₁₋₈alkyl, phenyl or phenyloxy, optionally substituted with halogen, hydroxy, C₁₋₄alkoxy, C₁₋₄alkyl, C₆₋₁₀aryl, carboxy, cyano, nitro, sulfonamido, sulfonate, phosphate, sulfonic acid, amino or substituted amino, wherein the amino is singly or doubly substituted by a C₁₋₄ alkyl and when doubly substituted, the alkyl groups is optionally linked to form a heterocycle; C₇₋₁₂arylalkyl; C₁₋₄alkylamino, phenylamino, C₇₋₁₂arylalkylamino, C₁₋₄alkoxy, or C₇₋₁₂arylalkyloxy; C₁₋₄alkylthio, phenylthio, C₇₋₁₂arylalkylthio, or -A(C₁₋₆alkyl)CONH(C₁₋₆alkyl)B- wherein A and B are independently

amino, mercapto, hydroxy or carboxyl;

provided that $R_1 \neq H$, when X is oxygen, methylene, or difluoromethylene, Y is OH, B is adenine, R_2 is absent, and R_3 is hydrogen;

provided that $R_1 \neq H$, when $n + m = 2$, X is oxygen, Y is OH, B is adenine, R_2 is absent, and R_3 is bromo, or 6-aminohexyl;

provided that $R_1 \neq H$, when $n + m = 2$, X is oxygen, Y is H, B is adenine, R_2 is absent, and R_3 is H;

provided that $R_2 \neq O$, when $n + m = 2$, X is oxygen, Y is OH, $R_1 = R_3 = H$, and B is adenine, adenine 1-oxide, or 1,N⁶-ethenoadenine;

provided that R_1 and R_2 do not form a 5-membered fused imidazole ring, when $n + m = 2$, X is oxygen, Y is OH, R_3 is H, and B is adenine, adenine 1-oxide, or ethenoadenine.

3. The compound according to Claim 1 or 2, wherein the ribosyl moieties are in the D- configuration.

4. The compound according to Claim 1 or 2, wherein the ribosyl moieties are in the L- configuration.

5. A pharmaceutical composition comprising a compound of Formula IA or IB as described in Claim 1 or 2, or a pharmaceutically acceptable salt thereof together with a pharmaceutically acceptable carrier therefor.

6. A method of treating chronic obstructive pulmonary diseases in a mammal by administering an effective chronic obstructive pulmonary disease treatment amount of a compound of Formula IA or IB as described in Claim 1 or 2.

7. A method of treating sinusitis, otitis media or nasolacrimal duct obstruction in a mammal by administering an effective mucus secretion clearing amount of a compound of Formula IA or IB as described in Claim 1 or 2.

8. A method of treating dry eye in a mammal by administering an effective dry eye treatment amount of a compound of Formula IA or IB as described in Claim 1 or 2.

5 9. A method of treating retinal detachment in a mammal by administering an effective retinal detachment treatment amount of a compound of Formula I as described in Claim 1 or 2.

10 10. A method of facilitating sputum induction in a mammal by administering an amount of a compound of Formula IA or IB as described in Claim 1 or 2, effective to facilitate sputum induction.

11. A method of facilitating expectoration in a mammal by administering an amount of a compound of Formula IA or IB as described in Claim 1 or 2, effective to facilitating expectoration.